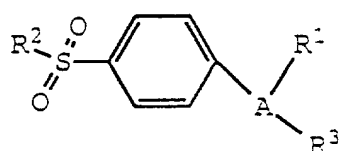


What is claimed is :

1. A method to suppress immune, acute or delayed-type hypersensitivity response in a subject, said
 5 method comprising treating the subject with a therapeutically-effective amount of a 5-lipoxygenase inhibitor and a cyclooxygenase-2 inhibitor selected from Dupont Dup 697, Taisho NS-398, meloxicam, flosulide and compounds of Formula I



I

15 wherein A is a 5- or 6-member ring substituent selected from partially unsaturated or unsaturated heterocyclo and carbocyclic rings;

20 wherein R¹ is at least one substituent selected from heterocyclo, cycloalkyl, cycloalkenyl and aryl, wherein R¹ is optionally substituted at a substitutable position with one or more radicals selected from alkyl, haloalkyl, cyano, carboxyl, alkoxycarbonyl, hydroxyl, hydroxyalkyl, haloalkoxy, amino, alkylamino, arylamino, nitro, alkoxyalkyl, alkylsulfinyl, halo, alkoxy and alkylthio;

25 wherein R² is selected from alkyl, and amino; and

30 wherein R³ is a radical selected from halo, alkyl, alkenyl, alkynyl, oxo, cyano, carboxyl, cyanoalkyl, heterocycloxy, alkyloxy, alkylthio, alkylcarbonyl, cycloalkyl, aryl, haloalkyl, heterocyclo, cycloalkenyl, aralkyl, heterocycloalkyl, acyl, alkylthioalkyl, hydroxyalkyl, alkoxycarbonyl, arylcarbonyl, aralkylcarbonyl, aralkenyl, alkoxyalkyl, arylthioalkyl, aryloxyalkyl, aralkylthioalkyl, aralkoxyalkyl, alkoxyaralkoxyalkyl, alkoxycarbonylalkyl, aminocarbonyl, aminocarbonylalkyl, alkylaminocarbonyl, N-arylaminocarbonyl, N-alkyl-N-arylaminocarbonyl,

35

alkylaminocarbonylalkyl, carboxyalkyl, alkylamino, N-aryl amino, N-aralkyl amino, N-alkyl-N-aralkyl amino, N-alkyl-N-aryl amino, aminoalkyl, alkylaminoalkyl, N-aryl aminoalkyl, N-aralkyl aminoalkyl, N-alkyl-N-aralkyl aminoalkyl, N-alkyl-N-aryl aminoalkyl, aryloxy, aralkoxy, arylthio, aralkylthio, alkylsulfinyl, alkylsulfonyl, aminosulfonyl, alkylaminosulfonyl, N-aryl aminosulfonyl, arylsulfonyl, N-alkyl-N-aryl aminosulfonyl;

10 or a pharmaceutically-acceptable salt thereof.

2. The method of Claim 1 wherein said 5-lipoxygenase inhibitor and said cyclooxygenase-2 inhibitor are administered in a sequential manner.

3. The method of Claim 1 wherein said 5-lipoxygenase inhibitor and said cyclooxygenase-2 inhibitor are administered in a substantially simultaneous manner.

4. The method of Claim 1 wherein the 5-lipoxygenase inhibitor is selected from masoprocol, tenidap, zileuton, flobufen, lonapalene, tagorizine, Abbott A-121798, Abbott A-76745, N'-[[5-(4-fluorophenoxy)furan-2-yl]-1-methyl-2-propynyl]-N'-hydroxyurea (Abbott A-78773), (R)(+)N'-[[5-(4-fluorophenoxy)furan-2-yl]-1-methyl-2-propynyl]-N-hydroxyurea (Abbott A-79175), Abbott ABT 761, Dainippon AL-3264, Bayer Bay-x-1005, Biofor BF-389, bunaprolast, Cytomed CMI-392, Takeda CV-6504, Efamol EF-40, Ciba-Geigy CGS-26529, enazadrem phosphate, Leo Denmark ETH-615, flezelastine hydrochloride, lonapalene, Merck Frosst L 663536, Merck Frosst L 699333, Merckle ML-3000, 3M Pharmaceuticals R-840, rilopirox, Schering Plough SCH 40120, tepoxalin, linazolast (TMK-688), Tanabe T-757, Tanabe T-799, Zeneca ZD 7717, Zeneca ZM-216800, Zeneca ZM 230487, and Zeneca ZD-2138.

5. The method of Claim 4 wherein the 5-lipoxygenase inhibitor is selected from tenidap, zileuton, flobufen, lonapalene, tagorizine, Abbott A-121798, Abbott A-76745, N'-[[5-(4-fluorophenoxy)furan-2-yl]-1-methyl-2-propynyl]-N'-hydroxyurea (Abbott A-78773), (R)(+)N'-[[5-(4-fluorophenoxy)furan-2-yl]-1-methyl-2-propynyl]-N-hydroxyurea (Abbott A-79175), Abbott ABT 761, Ciba-Geigy CGS-26529, Biofor BF-389, Cytomed CMI-392, Leo Denmark ETH-615, lonapalene, Merck Frosst L 699333, Merckle ML-3000, 3M Pharmaceuticals R-840, linazolast (TMK-688), Tanabe T-757, Tanabe T-799, Zeneca ZD 7717, Zeneca ZM-216300, Zeneca ZM 230487, and Zeneca ZD-2138.

6. The method of Claim 1 wherein A is selected from oxazolyl, isoxazolyl, dihydrofuryl, imidazolyl, and pyrazolyl; wherein R¹ is selected from 5- and 6-membered heterocyclo, and aryl selected from phenyl, biphenyl and naphthyl, wherein R¹ is optionally substituted at a substitutable position with one or more radicals selected from lower alkyl, lower haloalkyl, cyano, carboxyl, lower alkoxycarbonyl, hydroxyl, lower hydroxyalkyl, lower haloalkoxy, amino, lower alkylamino, phenylamino, nitro, lower alkoxyalkyl, lower alkylsulfinyl, halo, lower alkoxy and lower alkylthio; wherein R² is amino; and wherein R³ is a radical selected from oxo, cyano, carboxyl, lower alkoxycarbonyl, lower carboxyalkyl, lower cyanoalkyl, halo, lower alkyl, lower alkyloxy, lower cycloalkyl, phenyl, lower haloalkyl, 5- or 6-membered heterocyclo, lower hydroxylalkyl, lower aralkyl, acyl, phenylcarbonyl, lower alkoxyalkyl, 5- or 6-membered heteroaryloxy, aminocarbonyl, lower alkylaminocarbonyl, lower alkylamino, lower aminoalkyl, lower alkylaminoalkyl, phenyloxy, and lower aralkoxy; or a pharmaceutically-acceptable salt thereof.

7. The method of Claim 6 wherein A is selected from oxazolyl, isoxazolyl, imidazolyl, and pyrazolyl; wherein R¹ is phenyl optionally substituted at a substitutable position with one or more radicals selected from methyl, ethyl, isopropyl, butyl, *tert*-butyl, isobutyl, pentyl, hexyl, trifluoromethyl, cyano, carboxyl, methoxycarbonyl, hydroxyl, hydroxymethyl, trifluoromethoxy, amino, N-methylamino, N,N-dimethylamino, N-ethylamino, N,N-dipropylamino, N-butylamino, N-methyl-N-ethylamino, nitro, methoxymethyl, methylsulfinyl, fluoro, chloro, bromo, methoxy, ethoxy, propoxy, *n*-butoxy, pentoxy, and methylthio; wherein R² is amino; and wherein R³ is a radical selected from oxo, cyano, carboxyl, methoxycarbonyl, ethoxycarbonyl, carboxypropyl, carboxymethyl, carboxyethyl, cyanomethyl, fluoro, chloro, bromo, methyl, ethyl, isopropyl, butyl, *tert*-butyl, isobutyl, pentyl, hexyl, fluoromethyl, difluoromethyl, trifluoromethyl, chloromethyl, dichloromethyl, trichloromethyl, pentafluoroethyl, heptafluoropropyl, fluoromethyl, difluoroethyl, difluoropropyl, dichloroethyl, dichloropropyl, methoxy, ethoxy, propoxy, *n*-butoxy, pentoxy, cyclohexyl, phenyl, pyridyl, thienyl, thiazolyl, oxazolyl, furyl, pyrazinyl, hydroxylmethyl, hydroxylpropyl, benzyl, formyl, phenylcarbonyl, methoxymethyl, furylmethoxy, aminocarbonyl, N-methylaminocarbonyl, N,N-dimethylaminocarbonyl, N,N-dimethylamino, N-ethylamino, N,N-dipropylamino, N-butylamino, N-methyl-N-ethylamino, aminomethyl, N,N-dimethylaminomethyl, N-methyl-N-ethylaminomethyl, benzyloxy, and phenyloxy; or a pharmaceutically-acceptable salt thereof.

8. The method of Claim 7 selected from compounds, their prodrugs and their pharmaceutically-acceptable salts, of the group consisting of

3-(3,4-difluorophenyl)-4-(4-methylsulfonylphenyl)-
2-(5H)-furanone;

3-phenyl-4-(4-methylsulfonylphenyl)-2-(5H)-furanone;

5 4-[5-(4-chlorophenyl)-3-(trifluoromethyl)-1H-
pyrazol-1-yl]benzenesulfonamide;

4-[5-(4-methylphenyl)-3-(trifluoromethyl)-1H-
pyrazol-1-yl]benzenesulfonamide;

10 4-[5-(3-fluoro-4-methoxyphenyl)-3-
(difluoromethyl)-1H-pyrazol-1-yl]benzenesulfonamide;

3-[1-[4-(methylsulfonyl)phenyl]-4-trifluoromethyl-
1H-imidazol-2-yl]pyridine;

2-methyl-5-[1-[4-(methylsulfonyl)phenyl]-4-
trifluoromethyl-1H-imidazol-2-yl]pyridine;

15 4-[2-(5-methylpyridin-3-yl)-4-(trifluoromethyl)-
1H-imidazol-1-yl]benzenesulfonamide;

4-[5-methyl-3-phenylisoxazol-4-
yl]benzenesulfonamide;

20 4-[5-hydroxyethyl-3-phenylisoxazol-4-
yl]benzenesulfonamide;

[2-trifluoromethyl-5-(3,4-difluorophenyl)-4-
oxazolyl]benzenesulfonamide;

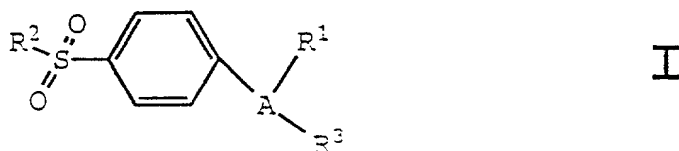
4-[2-methyl-4-phenyl-5-
oxazolyl]benzenesulfonamide; and

25 4-[5-(3-fluoro-4-methoxyphenyl)-2-trifluoromethyl)-
4-oxazolyl]benzenesulfonamide.

9. A combination comprising a therapeutically-
effective amount of a cyclooxygenase-2 inhibitor, a 5-
30 lipxygenase inhibitor and an immunosuppressive drug
selected from antiproliferative agents,
antiinflammatory-acting compounds and inhibitors of
leukocyte activation.

35 10. The combination of Claim 9 wherein the
cyclooxygenase-2 inhibitor is selected from Dupont Dup-

697, Taisho NS-398, meloxicam, flosulide and compounds of Formula I



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wherein A is a 5- or 6-member ring substituent selected from partially unsaturated or unsaturated heterocyclo and carbocyclic rings;

wherein R¹ is at least one substituent selected from heterocyclo, cycloalkyl, cycloalkenyl and aryl, wherein R¹ is optionally substituted at a substitutable position with one or more radicals selected from alkyl, haloalkyl, cyano, carboxyl, alkoxycarbonyl, hydroxyl, hydroxyalkyl, haloalkoxy, amino, alkylamino, arylamino, nitro, alkoxyalkyl, alkylsulfinyl, halo, alkoxy and alkylthio;

wherein R² is selected from alkyl, and amino; and

wherein R³ is a radical selected from halo, alkyl, alkenyl, alkynyl, oxo, cyano, carboxyl, cyanoalkyl, heterocycloxy, alkyloxy, alkylthio, alkylcarbonyl, cycloalkyl, aryl, haloalkyl, heterocyclo, cycloalkenyl, aralkyl, heterocycloalkyl, acyl, alkylthioalkyl, hydroxyalkyl, alkoxycarbonyl, arylcarbonyl, aralkylcarbonyl, aralkenyl, alkoxyalkyl, arylthioalkyl, aryloxyalkyl, aralkylthioalkyl, aralkoxyalkyl, alkoxyaralkoxyalkyl, alkoxycarbonylalkyl, aminocarbonyl, aminocarbonylalkyl, alkylaminocarbonyl, N-arylaminocarbonyl, N-alkyl-N-arylaminocarbonyl, alkylaminocarbonylalkyl, carboxyalkyl, alkylamino, N-arylamino, N-aralkylamino, N-alkyl-N-aralkylamino, N-alkyl-N-arylamino, aminoalkyl, alkylaminoalkyl, N-arylaminalkyl, N-aralkylaminalkyl, N-alkyl-N-aralkylaminalkyl, N-alkyl-N-arylaminalkyl, aryloxy, aralkoxy, arylthio, aralkylthio, alkylsulfinyl, alkylsulfonyl, aminosulfonyl, alkylaminosulfonyl, N-

arylaminosulfonyl, arylsulfonyl, N-alkyl-N-arylaminosulfonyl;

or a pharmaceutically-acceptable salt thereof.

- 5 11. The combination of Claim 9 wherein the 5-lipoxygenase inhibitor is selected from masoprocol, tenidap, zileuton, flubufen, lonapalene, tagorizine, Abbott A-121798, Abbott A-76745, N'-[[5-(4-fluorophenoxy)furan-2-yl]-1-methyl-2-propynyl]-N'-
- 10 hydroxyurea (Abbott A-78773), (R)(+)N'-[[5-(4-fluorophenoxy)furan-2-yl]-1-methyl-2-propynyl]-N-hydroxyurea (Abbott A-79175), Abbott ABT 761, Dainippon AL-3264, Bayer Bay-x-1005, Biofor BF-389, bunaprolast, Cytomed CMI-392, Takeda CV-6504, Efamol EF-40,
- 15 enazadrem phosphate, Leo Denmark ETH-615, flezelastine hydrochloride, Merck Frosst L 663536, Merckle ML-3000, 3M Pharmaceuticals R-840, rilopirox, Schering Plough SCH 40120, tepoxalin, linazolast (TMK-688), Tanabe T-757, Tanabe T-799, Zeneca ZD-2138, Abbott A 72694,
- 20 Abbott A-80263, Biofor BF-397, Bristol-Myers Squibb BU-4601A, carbazoycin C, lagunamycin, Wellcome BW-70C, Ciba-Geigy CGS-26529, Warner-Lambert CI 1004, Warner-Lambert PD-136005, Warner-Lambert PD-145246, Eisai E 3040, Fujirebio F-1322, Fisons FPL-64170, Fujisawa FR 110302, Nippon Hypox HX 0386, Merck & Co L-699333,
- 25 Merck Frosst L 739010, Lilly LY-269415, Lilly LY 178002, Meiji Milk MM-7002, Hoechst Roussel P 8892, Hoechst Roussel P 8977, SmithKline Beecham SB-202235, Green Cross SS-81-OH, Terumo Keio University TMK 685,
- 30 American Home Products WAY-121520, American Home Products WAY-125007, Zeneca ZD 7717, Zeneca ZM-216800, Zeneca ZM 230487, 1,2-dihydro-n-(2-thiazolyl)-1-oxopyrrolo(3,2,1-kl)phenothiazine-1-carboxamide, Abbott A-65260, Abbott A-69412, Abbott Abbott-63162, American
- 35 Home Products AHR-5333, Bayer Bay-q-1531, Boehringer Ingelheim BI-L-357, Boehringer Ingelheim BI-L-93BS, Boehringer Ingelheim BIL 226XX, Bristol-Myers Squibb

BMY-30094, carbazomycin B, Wellcome BW 4C, Wellcome BW-
 B218C, Wellcome BW-B70C, Chauvin CBS-1114, Ciba-Geigy
 CGS-21595, Ciba-Geigy CGS-22745, Ciba-Geigy CGS-23885,
 Ciba-Geigy CGS 24891, Ciba-Geigy CGS-8515, Chiesi CHF-
 5 1909, Warner-Lambert CI-986, Warner-Lambert CI 987,
 cirsilinol, docebenone, DuPont Merck DuP-654, Eisai E
 5110, Eisai E-6080, Green Cross EN-105, enofelast,
 epocarbazolin-A, eprovafen, evandamine, forsythiaside,
 Fisons FPL 62064, Glaxo GR-80907, Zeneca ICI-211965,
 10 isoflavans, Kyowa Hakko KF-8940, Merck & Co L-651392,
 Merck & Co L-651896, Merck & Co L-652343, Merck & Co
 L-656224, Merck & Co L-670630, Merck & Co L-674636,
 Merck & Co L-691816, Lilly LY-233569, Lilly LY-280810,
 Merck & Co MK-591, Merck & Co MK-886, nitrosoxacin-A,
 15 Ono ONO-5349, Ono ONO-LP-219, Ono ONO-LP-269, Warner-
 Lambert PD-127443, Purdue Frederick PF-5901, Sandoz QA-
 208-199, Johnson & Johnson R-68151, Johnson & Johnson
 R-85355, Rhone-Poulenc Rorer Rev-5367, Rhone-Poulenc
 Rorer RG-5901-A, Rhone-Poulenc Rorer RG-6866, Roussel-
 20 Uclaf RU-46057, Searle SC-41661A, Searle SC-45662,
 Sandoz SDZ-210-610, SmithKline Beecham SK&F-104351,
 SmithKline Beecham SK&F-104493, SmithKline Beecham
 SK&F-105809, Synthelabo SL-81-0433, Teijin TEI-8005,
 Terumo TMK-777, Terumo TMK-781, Terumo TMK-789, Terumo
 25 TMK-919, Terumo TMK-992, Teikoku Hormone TZI-2721,
 Teikoku Hormone TZI-41127, American Home Products WAY-
 120739, American Home Products WY 47288, American Home
 Products Wy-48252, American Home Products Wy-50295, and
 Yoshitomi Y-19432.

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12. The combination of Claim 11 wherein the 5-
 lipooxygenase inhibitor is selected from masoprocol,
 tenidap, zileuton, flubufen, lonapalene, tagorizine,
 Abbott A-121798, Abbott A-76745, N'-[[5-(4-
 35 fluorophenoxy)furan-2-yl]-1-methyl-2-propynyl]-N'-
 hydroxyurea (Abbott A-78773), (R)(+)N'-[[5-(4-
 fluorophenoxy)furan-2-yl]-1-methyl-2-propynyl]-N-

hydroxyurea (Abbott A-79175), Abbott ABT 761, Dainippon AL-3264, Bayer Bay-x-1005, Biofor BF-389, bunaprolast, Cytomed CMI-392, Takeda CV-6504, Efamol EF-40, Ciba-Geigy CGS-26529, enazadrem phosphate, Leo Denmark ETH-5 615, flezelastine hydrochloride, lonapalene, Merck Frosst L 663536, Merck Frosst L 699333, Merckle ML-3000, 3M Pharmaceuticals R-840, rilopirox, Schering Plough SCH 40120, tepoxalin, linazolast (TMK-688), Tanabe T-757, Tanabe T-799, Zeneca ZD 7717, Zeneca ZM-10 216800, Zeneca ZM 230487, and Zeneca ZD-2138.

13. The combination of Claim 12 wherein the 5-lipoxygenase inhibitor is selected from tenidap, zileuton, flobufen, lonapalene, tagorizine, Abbott A-15 121798, Abbott A-76745, N'-[[5-(4-fluorophenoxy)furan-2-yl]-1-methyl-2-propynyl]-N'-hydroxyurea (Abbott A-78773), (R)(+)N'-[[5-(4-fluorophenoxy)furan-2-yl]-1-methyl-2-propynyl]-N-hydroxyurea (Abbott A-79175), Abbott ABT 761, Ciba-Geigy CGS-26529, Biofor BF-389, 20 Cytomed CMI-392, Leo Denmark ETH-615, lonapalene, Merck Frosst L 699333, Merckle ML-3000, 3M Pharmaceuticals R-840, linazolast (TMK-688), Tanabe T-757, Tanabe T-799, Zeneca ZD 7717, Zeneca ZM-216800, Zeneca ZM 230487, and Zeneca ZD-2138.

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14. The combination of Claim 10 wherein A is selected from oxazolyl, isoxazolyl, thienyl, dihydrofuryl, furyl, pyrrolyl, pyrazolyl, thiazolyl, imidazolyl, isothiazolyl, cyclopentenyl, phenyl, and 30 pyridyl; wherein R¹ is selected from 5- and 6-membered heterocyclo, lower cycloalkyl, lower cycloalkenyl and aryl selected from phenyl, biphenyl and naphthyl, wherein R¹ is optionally substituted at a substitutable position with one or more radicals selected from lower 35 alkyl, lower haloalkyl, cyano, carboxyl, lower alkoxycarbonyl, hydroxyl, lower hydroxyalkyl, lower haloalkoxy, amino, lower alkylamino, phenylamino,

nitro, lower alkoxyalkyl, lower alkylsulfinyl, halo, lower alkoxy and lower alkylthio; wherein R^2 is selected from lower alkyl and amino; and wherein R^3 is a radical selected from halo, lower alkyl, oxo, cyano, carboxyl, lower cyanoalkyl, heteroaryloxy, lower alkyloxy, lower cycloalkyl, phenyl, lower haloalkyl, 5- or 6-membered heterocyclo, lower hydroxylalkyl, lower aralkyl, acyl, phenylcarbonyl, lower alkoxyalkyl, heteroaryloxy, alkoxycarbonyl, aminocarbonyl, alkylaminocarbonyl, alkylamino, aminoalkyl, alkylaminoalkyl, aryloxy, and aralkoxy; or a pharmaceutically-acceptable salt thereof.

15. The combination of Claim 14 wherein A is selected from oxazolyl, isoxazolyl, dihydrofuryl, imidazolyl, and pyrazolyl; wherein R^1 is selected from 5- and 6-membered heterocyclo, and aryl selected from phenyl, biphenyl and naphthyl, wherein R^1 is optionally substituted at a substitutable position with one or more radicals selected from lower alkyl, lower haloalkyl, cyano, carboxyl, lower alkoxycarbonyl, hydroxyl, lower hydroxyalkyl, lower haloalkoxy, amino, lower alkylamino, phenylamino, nitro, lower alkoxyalkyl, lower alkylsulfinyl, halo, lower alkoxy and lower alkylthio; wherein R^2 is amino; and wherein R^3 is a radical selected from oxo, cyano, carboxyl, lower alkoxycarbonyl, lower carboxyalkyl, lower cyanoalkyl, halo, lower alkyl, lower alkyloxy, lower cycloalkyl, phenyl, lower haloalkyl, 5- or 6-membered heterocyclo, lower hydroxylalkyl, lower aralkyl, acyl, phenylcarbonyl, lower alkoxyalkyl, 5- or 6-membered heteroaryloxy, aminocarbonyl, lower alkylaminocarbonyl, lower alkylamino, lower aminoalkyl, lower alkylaminoalkyl, phenyloxy, and lower aralkoxy; or a pharmaceutically-acceptable salt thereof.

16. The combination of Claim 15 wherein A is selected from oxazolyl, isoxazolyl, imidazolyl, and pyrazolyl; wherein R¹ is phenyl optionally substituted at a substitutable position with one or more radicals selected from methyl, ethyl, isopropyl, butyl, tert-butyl, isobutyl, pentyl, hexyl, trifluoromethyl, cyano, carboxyl, methoxycarbonyl, hydroxyl, hydroxymethyl, trifluoromethoxy, amino, N-methylamino, N,N-dimethylamino, N-ethylamino, N,N-dipropylamino, N-butylamino, N-methyl-N-ethylamino, nitro, methoxymethyl, methylsulfinyl, fluoro, chloro, bromo, methoxy, ethoxy, propoxy, n-butoxy, pentoxy, and methylthio; wherein R² is amino; and wherein R³ is a radical selected from oxo, cyano, carboxyl, methoxycarbonyl, ethoxycarbonyl, carboxypropyl, carboxymethyl, carboxyethyl, cyanomethyl, fluoro, chloro, bromo, methyl, ethyl, isopropyl, butyl, tert-butyl, isobutyl, pentyl, hexyl, fluoromethyl, difluoromethyl, trifluoromethyl, chloromethyl, dichloromethyl, trichloromethyl, pentafluoroethyl, heptafluoropropyl, fluoromethyl, difluoroethyl, difluoropropyl, dichloroethyl, dichloropropyl, methoxy, ethoxy, propoxy, n-butoxy, pentoxy, cyclohexyl, phenyl, pyridyl, thienyl, thiazolyl, oxazolyl, furyl, pyrazinyl, hydroxylmethyl, hydroxylpropyl, benzyl, formyl, phenylcarbonyl, methoxymethyl, furylmethoxy, aminocarbonyl, N-methylaminocarbonyl, N,N-dimethylaminocarbonyl, N,N-dimethylamino, N-ethylamino, N,N-dipropylamino, N-butylamino, N-methyl-N-ethylamino, aminomethyl, N,N-dimethylaminomethyl, N-methyl-N-ethylaminomethyl, benzyloxy, and phenyloxy; or a pharmaceutically-acceptable salt thereof.

17. The combination of Claim 16 selected from compounds, their prodrugs and their pharmaceutically-acceptable salts, of the group consisting of

3-(3,4-difluorophenyl)-4-(4-methylsulfonylphenyl)-
2-(5H)-furanone;

3-phenyl-4-(4-methylsulfonylphenyl)-2-(5H)-furanone;

4-[5-(4-chlorophenyl)-3-(trifluoromethyl)-1H-
5 pyrazol-1-yl]benzenesulfonamide;

4-[5-(4-methylphenyl)-3-(trifluoromethyl)-1H-
pyrazol-1-yl]benzenesulfonamide;

4-[5-(3-fluoro-4-methoxyphenyl)-3-
(difluoromethyl)-1H-pyrazol-1-yl]benzenesulfonamide;

10 3-[1-[4-(methylsulfonyl)phenyl]-4-trifluoromethyl-
1H-imidazol-2-yl]pyridine;

2-methyl-5-[1-[4-(methylsulfonyl)phenyl]-4-
trifluoromethyl-1H-imidazol-2-yl]pyridine;

4-[2-(5-methylpyridin-3-yl)-4-(trifluoromethyl)-
15 1H-imidazol-1-yl]benzenesulfonamide;

4-[5-methyl-3-phenylisoxazol-4-
yl]benzenesulfonamide;

4-[5-hydroxyethyl-3-phenylisoxazol-4-
yl]benzenesulfonamide;

20 [2-trifluoromethyl-5-(3,4-difluorophenyl)-4-
oxazolyl]benzenesulfonamide;

4-[2-methyl-4-phenyl-5-
oxazolyl]benzenesulfonamide; and

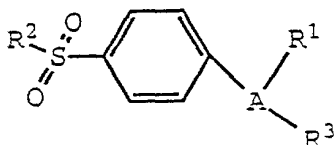
4-[5-(3-fluoro-4-methoxyphenyl)-2-trifluoromethyl)-
25 4-oxazolyl]benzenesulfonamide.

18. The composition of Claim 9 wherein the
leukocyte activation inhibitor is a cyclosporin.

30 19. The composition of Claim 18 wherein the
cyclosporin is cyclosporin A.

20. A pharmaceutical composition comprising a
pharmaceutically-acceptable carrier and a
35 therapeutically-effective amount of a 5-lipoxygenase
inhibitor, a cyclosporin and a cyclooxygenase-2

inhibitor selected from Dupont Dup 697, Taisho NS-398, meloxicam, flosulide and compounds of Formula I



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wherein A is a 5- or 6-member ring substituent selected from partially unsaturated or unsaturated heterocyclo and carbocyclic rings;

wherein R¹ is at least one substituent selected from heterocyclo, cycloalkyl, cycloalkenyl and aryl, wherein R¹ is optionally substituted at a substitutable position with one or more radicals selected from alkyl, haloalkyl, cyano, carboxyl, alkoxycarbonyl, hydroxyl, hydroxyalkyl, haloalkoxy, amino, alkylamino, arylamino, nitro, alkoxyalkyl, alkylsulfinyl, halo, alkoxy and alkylthio;

wherein R² is selected from alkyl, and amino; and

wherein R³ is a radical selected from halo, alkyl, alkenyl, alkynyl, oxo, cyano, carboxyl, cyanoalkyl, heterocyclooxy, alkyloxy, alkylthio, alkylcarbonyl, cycloalkyl, aryl, haloalkyl, heterocyclo, cycloalkenyl, aralkyl, heterocycloalkyl, acyl, alkylthioalkyl, hydroxyalkyl, alkoxycarbonyl, arylcarbonyl, aralkylcarbonyl, aralkenyl, alkoxyalkyl, arylthioalkyl, aryloxyalkyl, aralkylthioalkyl, aralkoxyalkyl, alkoxyaralkoxyalkyl, alkoxycarbonylalkyl, aminocarbonyl, aminocarbonylalkyl, alkylaminocarbonyl, N-arylaminocarbonyl, N-alkyl-N-arylaminocarbonyl, alkylaminocarbonylalkyl, carboxyalkyl, alkylamino, N-arylamino, N-aralkylamino, N-alkyl-N-aralkylamino, N-alkyl-N-arylamino, aminoalkyl, alkylaminoalkyl, N-arylaminalkyl, N-aralkylaminalkyl, N-alkyl-N-aralkylaminalkyl, N-alkyl-N-arylaminalkyl, aryloxy, aralkoxy, arylthio, aralkylthio, alkylsulfinyl, alkylsulfonyl, aminosulfonyl, alkylaminosulfonyl, N-

arylaminosulfonyl, arylsulfonyl, N-alkyl-N-arylaminosulfonyl;

or a pharmaceutically-acceptable salt thereof.

- 5 21. The method of Claim 1 further characterized
by suppressing immune response in a subject susceptible
to or afflicted with rejection of an organ transplanted
to said subject; graft versus host disease; an
autoimmune disease, an inflammatory disease, or a
10 condition with underlying autoimmune or inflammatory
reactivities or responses; an allergy; asthma; airway
hypersensitivity; septic shock; myesthemia gravis;
autoimmune thyroiditis; Grave's disease; autoimmune
hemolytic anemia; autoimmune thromboeytopenia purpura;
15 mixed connective tissue disease; idiopathic Addison's
disease; Sjogren's syndrome; urticaria; an acute
hypersensitivity response or a delayed hypersensitivity
response; Goodpasture's syndrome; hemolytic anemia;
contact dermatitis; granuloma; antibody-induced
20 thrombocytopenia; hypersensitivity pneumonitis;
glomerulonephritis; thyroiditis; encephalomyelitis; or
meningitis.